In re application of : Annapragada et al. Page 2 of 17

Application No. : 10/830,190 Filing Date : April 21, 2004

Examiner : Perreira, Melissa Jean

Title : Compositions and methods for enhancing contrast in

imaging

AMENDMENTS TO THE CLAIMS

1. (Currently amended) A composition for enhancing contrast of one or more areas of a subject for X-ray imaging when administered to the subject, the composition comprising:

liposomes, each liposome the liposomes encapsulating one or more iodinated nonradioactive contrast-enhancing agents, and each liposome the liposomes comprising: cholesterol, at least one phospholipid, and at least one phospholipid which is derivatized with a polymer chain,

wherein the average diameter of the liposomes is less than 150 nanometers.

- 2. (Previously amended) The composition of claim 1, wherein the X-ray imaging is computed tomography.
- 3. (Currently amended) The composition of claim 1, wherein the <u>iodinated</u> nonradioactive contrast-enhancing agents are selected from at least one of: iodinated ionic compounds, iodinated nonionic compounds, and mixtures <u>thereof</u>.
- 4. (Previously amended) The composition of claim 3, wherein a suspension of the liposomes has a concentration of at least 30 milligrams of iodine per milliliter of the suspension.
- 5. (Canceled).
- 6. (Previously amended) The composition of claim 1, wherein the average diameter of the liposomes is less than 120 nanometers.

In re application of : Annapragada et al. Page 3 of 17

Application No. : 10/830,190 Filing Date : April 21, 2004

Examiner : Perreira, Melissa Jean

Title : Compositions and methods for enhancing contrast in

imaging

7. (Previously amended) The composition of claim 1, wherein the composition is capable of being administered to the bloodstream of the subject.

- 8. (Previously amended) The composition of claim 7, wherein the composition provides an enhanced contrast that remains detectable at least 30 minutes after administration.
- 9. (Previously amended) The composition of claim 7, wherein the composition provides an enhanced contrast of at least 50 Hounsfield units in at least part of at least one of a vasculature and an organ of the subject.
- 10. (Previously amended) The composition of claim 1, wherein the liposomes are PEGylated liposomes.
- 11. (Previously amended) The composition of claim 1, wherein the liposomes are targeted liposomes.
- 12.-24. (Canceled).
- 25. (Currently amended) A composition for enhancing contrast of one or more areas of a subject for X-ray imaging when administered to the subject, the composition comprising liposomes, each liposome the liposomes comprising:
 - at least one first lipid or phospholipid;
 - at least one second lipid or phospholipid which is derivatized with one or more polymers; and

In re application of : Annapragada et al. Page 4 of 17

Application No. : 10/830,190 Filing Date : April 21, 2004

Examiner : Perreira, Melissa Jean

Title : Compositions and methods for enhancing contrast in

imaging

at least one sterically bulky excipient capable of stabilizing the liposomes;

wherein the average diameter of the liposomes is less than 150 nanometers, and wherein each liposome encapsulates the liposomes encapsulate at least one iodinated nonradioactive contrast enhancing agent.

- 26. (Previously amended) The composition of claim 25, wherein the at least one first lipid or phospholipid comprises 1,2-dipalmitoyl-sn-glycero-3-phosphocholine (DPPC).
- 27. (Currently amended) The composition of claim 25, wherein the at least one second lipid or phospholipid which is derivatized with one or more polymers comprises [N-(carbonylmethoxypolyethyleneglycol 2000)-1,2-distearoyl-sn-glycero-3-phosphatidyleholine] N-carbamylmethoxypoly(ethylene glycol)-1,2-distearoyl-sn-glycerol-3-phosphoethanolamine (DSPE-MPEG2000).
- 28. (Previously amended) The composition of claim 25, wherein the at least one sterically bulky excipient is selected from at least one of: sterols, fatty alcohols, fatty acids, and mixtures thereof.
- 29. (Previously amended) The composition of claim 25, wherein the at least one sterically bulky excipient is cholesterol.
- 30. (Previously amended) The composition of claim 25, wherein the liposomes are not autoclaved.

In re application of : Annapragada et al. Page 5 of 17

Application No. : 10/830,190 Filing Date : April 21, 2004

Examiner : Perreira, Melissa Jean

Title : Compositions and methods for enhancing contrast in

imaging

31. (Currently amended) The composition of claim 25, wherein the liposomes are contained in a suspension medium, at least some of the <u>iodinated nonradioactive</u> contrast enhancing agent that has not been encapsulated by the liposomes having been removed from the suspension medium.

- 32. (Previously presented) The composition of claim 25, wherein the at least one first lipid or phospholipid is present in the amount of about 55 to about 75 mol %; the at least one second lipid or phospholipid which is derivatized with one or more polymers is present in the amount of about 1 to about 20 mol %; and the at least one sterically bulky excipient is present in the amount of about 25 to about 40 mol %.
- 33. (Currently amended) The composition of claim 32, wherein the at least one first lipid or phospholipid is hydrogenated soy phosphatidylcholine; the at least one second lipid or phospholipid which is derivatized with one or more polymers is [N-(carbonylmethoxypolyethyleneglycol 2000)-1,2-distearoyl-sn-glycero-3-phosphatidylcholine] N-carbamylmethoxypoly(ethylene glycol)-1,2-distearoyl-sn-glycerol-3-phosphoethanolamine (DSPE-MPEG2000); and the at least one sterically bulky excipient is cholesterol.